



## SEQUENCE LISTING

<110> FRANZOSO, GUIDO  
DESMAELE, ENRICO  
ZAZZERONI, FRANCESCA  
PAPA, SALVATORE

<120> METHODS AND COMPOSITIONS FOR MODULATING APOPTOSIS

<130> 21459-94575

<140> 10/626,905

<141> 2003-07-25

<150> PCT/US02/31548

<151> 2002-10-02

<150> 10/263,330

<151> 2002-10-02

<150> 60/328,811

<151> 2001-10-12

<150> 60/326,492

<151> 2001-10-02

<160> 53

<170> PatentIn Ver. 3.2

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<212> DNA

<213> Homo sapiens

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                   20                  25                  30  
 Gln Asp Arg Leu Thr Val Gly Val Tyr Glu Ser Ala Lys Leu Met Asn  
           35                  40                  45  
 Val Asp Pro Asp Ser Val Val Leu Cys Leu Leu Ala Ile Asp Glu Glu  
           50                  55                  60  
 Glu Glu Asp Asp Ile Ala Leu Gln Ile His Phe Thr Leu Ile Gln Ser  
           65                  70                  75                  80  
 Phe Cys Cys Asp Asn Asp Ile Asn Ile Val Arg Val Ser Gly Asn Ala  
                   85                  90                  95  
 Arg Leu Ala Gln Leu Leu Gly Glu Pro Ala Glu Thr Gln Gly Thr Thr  
           100                  105                  110  
 Glu Ala Arg Asp Leu His Cys Leu Pro Phe Leu Gln Asn Pro His Thr  
           115                  120                  125  
 Asp Ala Trp Lys Ser His Gly Leu Val Glu Val Ala Ser Tyr Cys Glu  
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 tctggtcgca cggaaggtt tttttgcctc ttgggttcgt atctggactt gtactttgct 180  
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Gln Asp Arg Leu Thr Val Gly Val Tyr Glu Ala Ala Lys Leu Met Asn
      35              40              45

Val Asp Pro Asp Ser Val Val Leu Cys Leu Leu Ala Ile Asp Glu Glu
      50              55              60

Glu Glu Asp Asp Ile Ala Leu Gln Ile His Phe Thr Leu Ile Gln Ser
      65              70              75              80

Phe Cys Cys Asp Asn Asp Ile Asp Ile Val Arg Val Ser Gly Met Gln
      85              90              95

Arg Leu Ala Gln Leu Leu Gly Glu Pro Ala Glu Thr Leu Gly Thr Thr
      100             105             110

Glu Ala Arg Asp Leu His Cys Leu Leu Val Thr Asn Cys His Thr Asp
      115             120             125

Ser Trp Lys Ser Gln Gly Leu Val Glu Val Ala Ser Tyr Cys Glu Glu
      130             135             140

Ser Arg Gly Asn Asn Gln Trp Val Pro Tyr Ile Ser Leu Glu Glu Arg
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<210> 5  
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 <212> DNA  
 <213> Homo sapiens

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cggctggcac aggaggagga gcccgggcgg gcgaggggag gccggagagc gccagggcct 180
gagctgccgg agcgggcgct gtgagtga gtgcagaaagca ggcgcccgcg cgctagccgt 240

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cctggaggaa gtgctcagca aagccctgag tcagcgcacg atcactgtcg ggggtgtacga 420
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<212> PRT

<213> Mus musculus

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Asp Lys Val Gly Asp Ala Leu Glu Glu Val Leu Ser Lys Ala Leu Ser
      20              25              30

Gln Arg Thr Ile Thr Val Gly Val Tyr Glu Ala Ala Lys Leu Leu Asn
      35              40              45

Val Asp Pro Asp Asn Val Val Leu Cys Leu Leu Ala Ala Asp Glu Asp
      50              55              60

Asp Asp Arg Asp Val Ala Leu Gln Ile His Phe Thr Leu Ile Gln Ala
      65              70              75              80

Phe Cys Cys Glu Asn Asp Ile Asn Ile Leu Arg Val Ser Asn Pro Gly
      85              90              95

Arg Leu Ala Glu Leu Leu Leu Leu Glu Thr Asp Ala Gly Pro Ala Ala
      100             105             110

Ser Glu Gly Ala Glu Gln Pro Pro Asp Leu His Cys Val Leu Val Thr
      115             120             125

Asn Pro His Ser Ser Gln Trp Lys Asp Pro Ala Leu Ser Gln Leu Ile
      130             135             140

Cys Phe Cys Arg Glu Ser Arg Tyr Met Asp Gln Trp Val Pro Val Ile
      145             150             155             160

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Asn Leu Pro Glu Arg  
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gctcaacgta gaccccgata acgtggtact gtgacctgct gctgctgacg aagacgacga 360  
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gaagaaggaa gctgcgagaa aagagaaatc caaggcaaaa gggacaaaaa ctacaaagca 900  
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<212> PRT  
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Gln Arg Thr Ile Thr Val Gly Val Tyr Glu Ala Ala Lys Leu Leu Asn  
35 40 45  
Val Asp Pro Asp Asn Val Val Leu Cys Leu Leu Ala Ala Asp Glu Asp  
50 55 60  
Asp Asp Arg Asp Val Ala Leu Gln Ile His Phe Thr Leu Ile Arg Ala  
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Phe Cys Cys Glu Asn Asp Ile Asn Ile Leu Arg Val Ser Asn Pro Gly  
85 90 95

Arg Leu Ala Glu Leu Leu Leu Leu Glu Asn Asp Ala Gly Pro Ala Glu  
 100 105 110

Ser Gly Gly Ala Ala Gln Thr Pro Asp Leu His Cys Val Leu Val Thr  
 115 120 125

Asn Pro His Ser Ser Gln Trp Lys Asp Pro Ala Leu Ser Gln Leu Ile  
 130 135 140

Cys Phe Cys Arg Glu Ser Arg Tyr Met Asp Gln Trp Val Pro Val Ile  
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Asn Leu Pro Glu Arg  
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Ala Gln Arg Gln Gly Cys Leu Thr Ala Gly Val Tyr Glu Ser Ala Lys  
 35 40 45

Val Leu Asn Val Asp Pro Asp Asn Val Thr Phe Cys Val Leu Ala Ala  
 50 55 60

Gly Glu Glu Asp Glu Gly Asp Ile Ala Leu Gln Ile His Phe Thr Leu  
 65 70 75 80

Ile Gln Ala Phe Cys Cys Glu Asn Asp Ile Asp Ile Val Arg Val Gly  
 85 90 95

Asp Val Gln Arg Leu Ala Ala Ile Val Gly Ala Gly Glu Glu Ala Gly  
 100 105 110

Ala Pro Gly Asp Leu His Cys Ile Leu Ile Ser Asn Pro Asn Glu Asp  
 115 120 125

Ala Trp Lys Asp Pro Ala Leu Glu Lys Leu Ser Leu Phe Cys Glu Glu  
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 <211> 1084  
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 <213> Mus musculus

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 aaaa 1084

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Ala His Gly Gln Gly Cys Leu Thr Ala Gly Val Tyr Glu Ser Ala Lys  
                   35                  40                  45

Val Leu Asn Val Asp Pro Asp Asn Val Thr Phe Cys Val Leu Ala Ala  
                   50                  55                  60

Asp Glu Glu Asp Glu Gly Asp Ile Ala Leu Gln Ile His Phe Thr Leu  
                   65                  70                  75                  80

Ile Gln Ala Phe Cys Cys Glu Asn Asp Ile Asp Ile Val Arg Val Gly  
                   85                  90                  95

Asp Val Gln Arg Leu Ala Ala Ile Val Gly Ala Asp Glu Glu Gly Gly  
                   100                  105                  110

Ala Pro Gly Asp Leu His Cys Ile Leu Ile Ser Asn Pro Asn Glu Asp  
                   115                  120                  125

Thr Trp Lys Asp Pro Ala Leu Glu Lys Leu Ser Leu Phe Cys Glu Glu  
                   130                  135                  140

Ser Arg Ser Phe Asn Asp Trp Val Pro Ser Ile Thr Leu Pro Glu  
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<210> 13  
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 <212> DNA  
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<220>  
 <223> Description of Artificial Sequence: Primer

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<210> 14  
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 <212> DNA  
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 <223> Description of Artificial Sequence: Primer

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<210> 15  
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<220>  
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 <210> 17  
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 <212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Primer

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<210> 21

<211> 41

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Primer

<400> 21

ggatggatat ccgaaattaa tccaagaaga cagagatgaa c

41

<210> 22

<211> 39

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Primer

<400> 22

ggataacgcg tcaccgtcct caaacttacc aaacgttta

39

<210> 23

<211> 39

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Primer

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39

<210> 24

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<212> DNA

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<223> Description of Artificial Sequence: Primer

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38

<210> 25  
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<212> DNA  
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<220>  
<223> Description of Artificial Sequence: Primer

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<210> 26  
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<220>  
<223> Description of Artificial Sequence: Primer

<400> 26  
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<210> 27  
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<220>  
<223> Description of Artificial Sequence: Primer

<400> 27  
aatattctct cc 12

<210> 28  
<211> 10  
<212> DNA  
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<220>  
<223> Description of Artificial Sequence: Primer

<400> 28  
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<210> 29  
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<212> DNA  
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<220>  
<223> Description of Artificial Sequence: Primer

<400> 29  
atcgattcca 10

<210> 30  
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<220>  
 <223> Description of Artificial Sequence: Primer

<400> 30  
 ggaaaccccg 10

<210> 31  
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<220>  
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<400> 31  
 ggaaatattg 10

<210> 32  
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 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Description of Artificial Sequence: Primer

<400> 32  
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<210> 33  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Description of Artificial Sequence: Primer

<400> 33  
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<210> 34  
 <211> 27  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Description of Artificial Sequence: Primer

<400> 34  
gatctgaatt ctacttactc tcaagac

27

<210> 35  
<211> 2695  
<212> DNA  
<213> Mus musculus

<400> 35  
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ttttggtttt tcgagacagg gtttctctgt gtagccctgg ctgtcctgga actcactctg 300  
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<210> 36  
 <211> 10  
 <212> DNA  
 <213> Mus musculus

<400> 36  
 gggactctcc 10

<210> 37  
 <211> 16  
 <212> DNA  
 <213> Mus musculus

<400> 37  
 ctagggactc tccggg 16

<210> 38  
 <211> 10  
 <212> DNA  
 <213> Mus musculus

<400> 38  
 ggggattcca 10

<210> 39  
 <211> 16  
 <212> DNA  
 <213> Mus musculus

<400> 39  
 cgaggggatt ccagac 16

<210> 40  
 <211> 10  
 <212> DNA  
 <213> Mus musculus

<400> 40  
 ggaaaccccg 10

<210> 41  
 <211> 16  
 <212> DNA  
 <213> Mus musculus

<400> 41  
 gctggaaacc ccgcgc 16

<210> 42  
 <211> 5  
 <212> PRT  
 <213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Synthetic  
peptide

<400> 42  
Val Asp Val Ala Asp  
1

<210> 43  
<211> 4  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Synthetic  
peptide

<400> 43  
Asp Glu Val Asp  
1

<210> 44  
<211> 4  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Synthetic  
peptide

<400> 44  
Val Glu Ile Asp  
1

<210> 45  
<211> 4  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Synthetic  
peptide

<400> 45  
Ile Glu Thr Asp  
1

<210> 46  
<211> 4  
<212> PRT  
<213> Artificial Sequence

&lt;220&gt;

&lt;223&gt; Description of Artificial Sequence: Synthetic peptide

&lt;400&gt; 46

Leu Glu His Asp

1

&lt;210&gt; 47

&lt;211&gt; 27

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Description of Artificial Sequence: Synthetic oligonucleotide

&lt;400&gt; 47

cgccaccatg gagatggtga acaccat

27

&lt;210&gt; 48

&lt;211&gt; 33

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Description of Artificial Sequence: Synthetic oligonucleotide

&lt;400&gt; 48

gtacaagggt atggctatgt caatgggagg tag

33

&lt;210&gt; 49

&lt;211&gt; 1392

&lt;212&gt; DNA

&lt;213&gt; Homo sapiens

&lt;400&gt; 49

```

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```



```

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gaacacagct tcatcaagcg ctacgagacg ctggaggtgg acgtggcgtc ctgggttcaag 1260
gatgtcatgg cgaagacctg agtcaccgcg gactaacggc gttccttgag ccagccccac 1320
cttggcccct tcttcaggtt agcttgcttt ggccggcggc caaccctctt ggggggccag 1380
ggcattggcc cc                                     1392

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<210> 50

<211> 401

<212> PRT

<213> Homo sapiens

<400> 50

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Met Ala Ala Ser Ser Leu Glu Gln Lys Leu Ser Arg Leu Glu Ala Lys
  1              5              10              15

```

```

Leu Lys Gln Glu Asn Arg Glu Ala Arg Arg Arg Ile Asp Leu Asn Leu
          20              25              30

```

```

Asp Ile Ser Pro Gln Arg Pro Arg Pro Thr Leu Gln Leu Pro Leu Ala
          35              40              45

```

```

Asn Asp Gly Gly Ser Arg Ser Pro Ser Ser Glu Ser Ser Pro Gln His
          50              55              60

```

```

Pro Thr Pro Pro Ala Arg Pro Arg His Met Leu Gly Leu Pro Ser Thr
          65              70              75              80

```

```

Leu Phe Thr Pro Arg Ser Met Glu Ser Ile Glu Ile Asp His Lys Leu
          85              90              95

```

```

Gln Glu Ile Met Lys Gln Thr Gly Tyr Leu Thr Ile Gly Gly Gln Arg
          100             105             110

```

```

Tyr Gln Ala Glu Ile Asn Asp Leu Glu Asn Leu Gly Glu Met Gly Ser
          115             120             125

```

```

Gly Thr Cys Gly Pro Val Trp Lys Met Arg Phe Arg Lys Thr Gly His
          130             135             140

```

```

Val Ile Ala Val Lys Gln Met Arg Arg Ser Gly Asn Lys Glu Glu Asn
          145             150             155             160

```

```

Lys Arg Ile Leu Met Asp Leu Asp Val Val Leu Lys Ser His Asp Cys
          165             170             175

```

```

Pro Tyr Ile Val Gln Cys Phe Gly Thr Phe Ile Thr Asn Thr Asp Val
          180             185             190

```

```

Phe Ile Ala Met Glu Leu Met Gly Thr Cys Ala Glu Lys Leu Lys Lys
          195             200             205

```

```

Arg Met Gln Gly Pro Ile Pro Glu Arg Ile Leu Gly Lys Met Thr Val
          210             215             220

```

Ala Ile Val Lys Ala Leu Tyr Tyr Leu Lys Glu Lys His Gly Val Ile  
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His Arg Asp Val Lys Pro Ser Asn Ile Leu Leu Asp Glu Arg Gly Gln  
 245 250 255

Ile Lys Leu Cys Asp Phe Gly Ile Ser Gly Arg Leu Val Asp Ser Lys  
 260 265 270

Ala Lys Thr Arg Ser Ala Gly Cys Ala Ala Tyr Met Ala Pro Glu Arg  
 275 280 285

Ile Asp Pro Pro Asp Pro Thr Lys Pro Asp Tyr Asp Ile Arg Ala Asp  
 290 295 300

Val Trp Ser Leu Gly Ile Ser Leu Val Glu Leu Ala Thr Gly Gln Phe  
 305 310 315 320

Pro Tyr Lys Asn Cys Lys Thr Asp Phe Glu Val Leu Thr Lys Val Leu  
 325 330 335

Gln Glu Glu Pro Pro Leu Leu Pro Gly His Met Gly Phe Ser Gly Asp  
 340 345 350

Phe Gln Ser Phe Val Lys Asp Cys Leu Thr Lys Asp His Arg Lys Arg  
 355 360 365

Pro Lys Tyr Asn Lys Leu Leu Glu His Ser Phe Ile Lys Arg Tyr Glu  
 370 375 380

Thr Leu Glu Val Asp Val Ala Ser Trp Phe Lys Asp Val Met Ala Lys  
 385 390 395 400

Thr

<210> 51  
 <211> 2313  
 <212> DNA  
 <213> Mus musculus

<400> 51  
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 gctgtgctgc ctatatggct cccgagcgca tgcacctcc agatccacc aagcctgact 900  
 atgacatccg agctgatgtg tggagcctgg gcatctcact ggtggagctg gcaacaggac 960

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```

<210> 52

<211> 346

<212> PRT

<213> Mus musculus

<400> 52

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Met Leu Gly Leu Pro Ser Thr Leu Phe Thr Pro Arg Ser Met Glu Ser
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Ile Glu Ile Asp Gln Lys Leu Gln Glu Ile Met Lys Gln Thr Gly Tyr
      20              25              30

Leu Thr Ile Gly Gly Gln Arg Tyr Gln Ala Glu Ile Asn Asp Leu Glu
      35              40              45

Asn Leu Gly Glu Met Gly Ser Gly Thr Cys Gly Gln Val Trp Lys Met
      50              55              60

Arg Phe Arg Lys Thr Gly His Ile Ile Ala Val Lys Gln Met Arg Arg
      65              70              75              80

Ser Gly Asn Lys Glu Glu Asn Lys Arg Ile Leu Met Asp Leu Asp Val
      85              90              95

Val Leu Lys Ser His Asp Cys Pro Tyr Ile Val Gln Cys Phe Gly Thr
      100             105             110

Phe Ile Thr Asn Thr Asp Val Phe Ile Ala Met Glu Leu Met Gly Ile
      115             120             125

Cys Ala Glu Lys Leu Lys Lys Arg Met Gln Gly Pro Ile Pro Glu Arg
      130             135             140

```

Ile	Leu	Gly	Lys	Met	Thr	Val	Ala	Ile	Val	Lys	Ala	Leu	Tyr	Tyr	Leu
145					150					155					160
Lys	Glu	Lys	His	Gly	Val	Ile	His	Arg	Asp	Val	Lys	Pro	Ser	Asn	Ile
				165					170					175	
Leu	Leu	Asp	Glu	Arg	Gly	Gln	Ile	Lys	Leu	Cys	Asp	Phe	Gly	Ile	Ser
			180					185					190		
Gly	Arg	Leu	Val	Asp	Ser	Lys	Ala	Lys	Thr	Arg	Ser	Ala	Gly	Cys	Ala
		195					200					205			
Ala	Tyr	Met	Ala	Pro	Glu	Arg	Ile	Asp	Pro	Pro	Asp	Pro	Thr	Lys	Pro
	210					215					220				
Asp	Tyr	Asp	Ile	Arg	Ala	Asp	Val	Trp	Ser	Leu	Gly	Ile	Ser	Leu	Val
225					230					235					240
Glu	Leu	Ala	Thr	Gly	Gln	Phe	Pro	Tyr	Lys	Asn	Cys	Lys	Thr	Asp	Phe
				245					250					255	
Glu	Val	Leu	Thr	Lys	Val	Leu	Gln	Glu	Glu	Pro	Pro	Leu	Leu	Pro	Gly
			260					265						270	
His	Met	Gly	Phe	Ser	Gly	Asp	Phe	Gln	Ser	Phe	Val	Lys	Asp	Cys	Leu
		275					280					285			
Thr	Lys	Asp	His	Arg	Lys	Arg	Pro	Lys	Tyr	Asn	Lys	Leu	Leu	Glu	His
	290					295					300				
Ser	Phe	Ile	Lys	His	Tyr	Glu	Ile	Leu	Glu	Val	Asp	Val	Ala	Ser	Trp
305					310					315					320
Phe	Lys	Asp	Val	Met	Ala	Lys	Thr	Asp	Ser	Pro	Arg	Thr	Ser	Gly	Val
				325					330					335	
Leu	Ser	Gln	His	His	Leu	Pro	Phe	Phe	Arg						
			340					345							

&lt;210&gt; 53

&lt;211&gt; 6

&lt;212&gt; PRT

&lt;213&gt; Artificial Sequence

&lt;220&gt;

<223> Description of Artificial Sequence: Synthetic  
6X-His tag

&lt;400&gt; 53

His His His His His His

1

5